

## INTERNATIONAL SEARCH REPORT

ational Application No  
/BE2004/000124

A. CLASSIFICATION OF SUBJECT MATTER  
IPC 7 A61K31/519 A61K31/5377 A61K31/541 A61K45/06 C07D475/04  
C07D475/08 C07D475/00 A61P37/00 A61P37/02 A61P37/06  
A61P9/00 A61P25/00 A61P35/00

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)  
IPC 7 A61K C07D

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, CHEM ABS Data, PAJ, WPI Data, BIOSIS, EMBASE, MEDLINE, SCISEARCH, BEILSTEIN Data

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 00/39129 A (WAER MARK JOSEPH ALBERT ; HERDEWIJN PIET ANDRE MAURITS M (BE); LEUVEN) 6 July 2000 (2000-07-06) page 1, line 4 - page 2, line 10 page 2, line 34 - page 3, line 12 page 4, lines 3-7 page 7, lines 31-34 page 8, lines 6-29 page 10, lines 6-31 page 12, lines 15-26 page 17, line 30 - page 18, line 13 page 19, lines 8-20 page 19, line 34 - page 20, line 14 page 20, lines 20-23 compounds 1,2,3,6,13-16,21-63,65,66 claims 1-8,13-17  ----- -/--	1-16, 20-24

☒ Further documents are listed in the continuation of box C.☒ Patent family members are listed in annex.

## \* Special categories of cited documents:

- \*A\* document defining the general state of the art which is not considered to be of particular relevance
- \*E\* earlier document but published on or after the international filing date
- \*L\* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- \*O\* document referring to an oral disclosure, use, exhibition or other means
- \*P\* document published prior to the international filing date but later than the priority date claimed

- \*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- \*X\* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- \*Y\* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- \*G\* document member of the same patent family

Date of the actual completion of the international search

12 April 2005

Date of mailing of the international search report

27/04/2005

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2  
NL - 2280 HV Rijswijk  
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  
Fax: (+31-70) 340-3016

Authorized officer

Cielen, E

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## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

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X	MATTER HANS ET AL: "Structural requirements for inhibition of the neuronal nitric oxide synthase (NOS-I): 3D-QSAR analysis of 4-oxo- and 4-amino-pteridine-based inhibitors" JOURNAL OF MEDICINAL CHEMISTRY, vol. 45, no. 14, 4 July 2002 (2002-07-04), pages 2923-2941, XP002313348 ISSN: 0022-2623	1-7
A	abstract page 2924, column 2, paragraph 3 compounds 264,265,246,247,301,303,305,306,310,311,313,315 compounds 317,318 page 2938, column 1, paragraphs 5,7 page 2938, column 2, paragraph 1	8-14
X	GB 785 353 A (MERCK & CO INC) 30 October 1957 (1957-10-30) page 1, lines 22-43 page 2, lines 72-87 page 2, lines 120-130 claims 1,8,10	1,2
X	DATABASE CA 'Online! CHEMICAL ABSTRACTS SERVICE, COLUMBUS, OHIO, US; KALDRIKYAN, M. A. ET AL: "Pteridine derivatives. I. Synthesis of some substituted 6,7-diarylpteridines" XP002313350 retrieved from STN Database accession no. 1976:560035 abstract & ARMYANSKII KHIMICHESKII ZHURNAL , 29(4), 337-41 CODEN: AYKZAN; ISSN: 0515-9628, 1976,	1-6
X	YAO, QIZINENG ET AL: "Pteridines. Part CXIII. Protection of pteridines" HELVETICA CHIMICA ACTA , 86(1), 1-12 CODEN: HCACAV; ISSN: 0018-019X, 2003, XP008041327 compound 28	1

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A	WO 01/21619 A (PFLEIDERER WOLFGANG ; KOTSONIS PETER (DE); SCHMIDT HARALD (DE); FROEHL) 29 March 2001 (2001-03-29) page 1, lines 10-13 page 4, lines 19-25 page 5, line 1 - page 7, line 10 page 15, lines 11-30 page 16, lines 11-14 page 21, line 16 - page 23, line 27 claims 1-8	1-14, 20-22
A	----- NICOLAUS B J R: "Symbiotic Approach to Drug Design" DECISION MAKING IN DRUG RESEARCH, XX, XX, 1983, pages 173-186, XP002197412 the whole document	
T	----- EP 1 479 682 A (4 AZA BIOSCIENCE NV) 24 November 2004 (2004-11-24) page 2, paragraph 1-3 page 24, paragraphs 67,70,71 page 25, paragraphs 73,74 page 27, paragraphs 87,88	8-17, 20-25
X	----- ISRAEL, MERVYN ET AL: "Pyrimidine derivatives. VII. Some condensed derivatives of 2,4,5-triamino-6-methylthiopyrimidine" JOURNAL OF PHARMACEUTICAL SCIENCES , 54(11), 1626-32 CODEN: JPMSAE; ISSN: 0022-3549, 1965, XP008045079 compounds XIIIA,XIIIB,XIV page 1631, column 1, paragraph 4 - column 2, paragraph 1	1,2
X	----- DATABASE BEILSTEIN 29 November 1988 (1988-11-29), XP002324247 abstract; compound BRN1184281	1,2
X	----- ELLIOTT R D ET AL: "Synthesis of N-10-methyl-4-thiofolic acid and related compounds." JOURNAL OF MEDICINAL CHEMISTRY. MAY 1975, vol. 18, no. 5, May 1975 (1975-05), pages 492-496, XP002324245 ISSN: 0022-2623 compounds 10,17-19 page 495, column 1, paragraph 2 page 495, column 1, last paragraph - column 2, paragraph 3	1
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## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

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A	US 3 159 628 A (PACHTER IRWIN J ET AL) 1 December 1964 (1964-12-01) column 4, lines 13-15 column 4, lines 36,37 claims 1,5	1-3,8-10
X	EP 0 134 922 A (DR. KARL THOMAE GMBH) 27 March 1985 (1985-03-27)  page 6, line 17 - page 7, line 4 examples 6,7,10,12,14,17,18 claims 1,9	1,2,5,6, 8,9,12, 13
X	EP 0 185 259 A (DR. KARL THOMAE GMBH) 25 June 1986 (1986-06-25)  page 2, lines 16-24 page 7, line 28 - page 8, line 12 page 14, line 8 - page 15, line 4 examples 2,3,6-8,12 page 22, lines 23-26 page 23, lines 5-20 examples 16,18-21,23,26-29,33 claims 1,2,7	1,2,5,6, 8,9,12, 13
X	FROEHLICH LOTHAR G ET AL: "Inhibition of neuronal nitric oxide synthase by 4-amino pteridine derivatives: Structure-activity relationship of antagonists of (6R)-5,6,7,8-tetrahydrobiopterin cofactor" JOURNAL OF MEDICINAL CHEMISTRY, vol. 42, no. 20, 7 October 1999 (1999-10-07), pages 4108-4121, XP002324246 ISSN: 0022-2623	1-7
A	abstract page 4108, column 2, paragraph 1 page 4109, column 1, paragraph 3 table 9 page 4113, column 2, paragraph 1 - page 4114, column 2, paragraph 1 page 4114, column 2, paragraph 3 page 4118, column 2, paragraph 5 - page 4119, column 1, paragraph 2	8-14
X	DE 19 21 308 A1 (C.H. BOEHRINGER SOHN) 7 January 1971 (1971-01-07) page 2, paragraph 1 page 13, lines 10,11 page 13, lines 1,2 page 13, line 15 - page 14, line 2  -/--	1,6,8,13

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Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 3 081 230 A (WEINSTOCK JOSEPH ET AL) 12 March 1963 (1963-03-12) column 1, lines 11-18 column 2, lines 3-27 example 24	1,5,8,12
X	DE 40 09 941 A1 (DR. KARL THOMAE GMBH, 7950 BIBERACH, DE) 2 October 1991 (1991-10-02) page 2, line 45 - page 3, line 15 page 3, lines 51,55 page 5, lines 25,26 claims 1,3,7,11	1,8,15, 16
X	WO 03/062240 A (FAUSTUS FORSCHUNGS CIE. TRANSLATIONAL CANCER RESEARCH GMBH; EISENBRAND) 31 July 2003 (2003-07-31) page 1, lines 5-10 page 2, line 20 - page 4, line 17 examples 3,6 claims 1,2,13	1,6,8,13
X	US 2 940 972 A (ROCH JOSEF) 14 June 1960 (1960-06-14) column 1, lines 15-39 column 3, lines 16-29 examples 3,7 columns 9-14; compounds 8,10,12,19,25,26,30,33,38,39 claims 1,4,7,8	1,6,8,13
X	DD 267 495 A1 (AKADEMIE DER WISSENSCHAFTEN DER DDR,DD) 3 May 1989 (1989-05-03) page 1, paragraph 5 - last paragraph	1,8
X	US 3 859 287 A (PARISH ET AL) 7 January 1975 (1975-01-07) column 1, lines 16-58 column 4, lines 30-35 column 4, line 66 - column 5, line 2 examples VII,XVIX,XX column 21, line 64 - column 22, line 5 figures 6,7	1,8
X	SPICKETT R G W ET AL: "THE SYNTHESIS OF COMPOUNDS WITH POTENTIAL ANTI-FOLIC ACID ACTIVITY. PART I. 7-AMINO- AND 7-HYDROXY-PTERIDINES" JOURNAL OF THE CHEMICAL SOCIETY, CHEMICAL SOCIETY. LETCHWORTH, GB, 1954, pages 2887-2891, XP008045137 ISSN: 0368-1769 compound III page 2892, paragraphs 4,5	1,4

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## C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	<p>WO 02/32507 A (ASTRAZENECA AB; BONNERT, ROGER; WALTERS, IAIN)  25 April 2002 (2002-04-25)  page 1, lines 3-5  page 1, line 31 - page 3, line 17  page 5, lines 6-15  page 6, lines 26,27  page 7, line 7 - page 8, line 29  page 9, lines 25-29  -----</p>	<p>1,8,20, 22</p>

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# INTERNATIONAL SEARCH REPORT

International application No.  
PCT/BE2004/000124

## Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:  
  
Although claims 20-25 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2. ☐ Claims Nos.:  
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. ☐ Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

see additional sheet

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☒ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:  
  
1-7 (entirely), 8-15 (partially), 16-17 (entirely), 20-23 (partially)  
24-25 (entirely)
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☒ No protest accompanied the payment of additional search fees.

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## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-4, 7-11, 14-17, 20-25 (all partially)

A pteridine derivative having the general formula as defined in claim 1, wherein X represents an oxygen atom.

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents an oxygen atom, as far as related to the prevention or treatment of immune and auto-immune disorders, and optionally further comprising one or more immuno-suppressants and/or immunomodulator drugs.

A method for the prevention or treatment of immune and auto-immune disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents an oxygen atom, and optionally further comprising one or more immuno-suppressants and/or immunomodulator drugs.

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2. claims: 1-4, 7-11, 14-17, 20-25 (all partially)

A pteridine derivative having the general formula as defined in claim 1, wherein X represents a group with the formula S(O)m.

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents a group with the formula S(O)m, as far as related to the prevention or treatment of immune and auto-immune disorders, and optionally further comprising one or more immuno-suppressants and/or immunomodulator drugs.

A method for the prevention or treatment of immune and auto-immune disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents a group with the formula S(O)m, and optionally further comprising one or more immuno-suppressants and/or immunomodulator drugs.

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3. claims: 1-4 (partially), 5-6 (entirely), 7-17 (partially), 20-25 (partially)



## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

A pteridine derivative having the general formula as defined in claim 1, wherein X represents a group with the formula NZ.

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents a group with the formula NZ, as far as related to the prevention or treatment of immune and auto-immune disorders, and optionally further comprising one or more immuno-suppressants and/or immunomodulator drugs.

A method for the prevention or treatment of immune and auto-immune disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents a group with the formula NZ, and optionally further comprising one or more immuno-suppressants and/or immunomodulator drugs.

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4. claims: 8-11, 14, 20-22 (all partially)

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents an oxygen atom, as far as related to the prevention or treatment of cardiovascular disorders.

A method for the prevention or treatment of cardiovascular disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents an oxygen atom.

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5. claims: 8-11, 14, 20-22 (all partially)

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents a group with the formula S(O)m, as far as related to the prevention or treatment of cardiovascular disorders.

A method for the prevention or treatment of cardiovascular disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents a group with the formula S(O)m.

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6. claims: 8-14, 20-22 (all partially)

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents a group with the formula NZ, as far as related to the prevention or treatment of cardiovascular disorders.

A method for the prevention or treatment of cardiovascular disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents a group with the formula NZ.

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## 7. claims: 8-11, 14, 20-22 (all partially)

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents an oxygen atom, as far as related to the prevention or treatment of disorders of the central nervous system.

A method for the prevention or treatment of disorders of the central nervous system comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents an oxygen atom.

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## 8. claims: 8-11, 14, 20-22 (all partially)

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents a group with the formula S(O)m, as far as related to the prevention or treatment of disorders of the central nervous system.

A method for the prevention or treatment of disorders of the central nervous system comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents a group with the formula S(O)m.

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## 9. claims: 8-14, 20-22 (all partially)

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents a group with the formula NZ, as far as related to the prevention or treatment of disorders of the central nervous system.

A method for the prevention or treatment of disorders of the central nervous system comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents a group with the formula NZ.

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## 10. claims: 8-11, 14-15, 18, 20-23, 26 (all partially)

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents an oxygen atom, as far as related to the prevention or treatment of cell proliferative disorders, and optionally further comprising one or more antineoplastic drugs.

A method for the prevention or treatment of cell proliferative disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents an oxygen atom, and optionally further comprising one or more antineoplastic drugs.

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## 11. claims: 8-11, 14-15, 18, 20-23, 26 (all partially)

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents a group with the formula S(O)m, as far as related to the prevention or treatment of cell proliferative disorders, and optionally further comprising one or more antineoplastic drugs.

A method for the prevention or treatment of cell proliferative disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents a group with the formula S(O)m, and optionally further comprising one or more antineoplastic drugs.

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## 12. claims: 8-15, 18, 20-23, 26 (all partially)

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8, wherein X represents a group with the formula NZ, as far as related to the prevention or treatment of cell proliferative disorders, and optionally further comprising one or more antineoplastic drugs.

A method for the prevention or treatment of cell proliferative disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20, wherein X represents a group with the formula NZ, and optionally further comprising one or more antineoplastic drugs.

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13. claims: 8-15 (partially), 19 (entirely), 20-23 (partially), 27 (entirely)

A pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 8 and further comprising one or more antiviral drugs.

A method for the prevention or treatment of antiviral disorders comprising the administration of a pharmaceutical composition comprising as an active principle at least one pteridine derivative having the general formula as defined in claim 20 and further comprising one or more antiviral drugs.

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## INTERNATIONAL SEARCH REPORT

Information on patent family members

national Application No

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